

Fig. IA

卷之三

Fig. 1B

152827360

1 GCTGTGGAA CCTCTCCACG CGCACGAAT CAGCCAACGA TTTCTGATAG ATTTTGGGA GTTGACCAAGG AGATGCAAGG GGTGAAGGAG CGCTTCCTAC
CGACACCCCTT GGAGGCTGC GCGTGTGTA GTCGGTGTCT AAGACTATC TAAAAACCTT CAAACTGGTC TCTACGTTCC CCACTCCCTC CGGAAGGATC
-40 MetLngL YvallysL GluArgLeuPro

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GCAATTCCCTT GAGACCCCTG TCTCGGGGG CCGGGGACT ACCGGCTCCG TCCCACGCTG GGTCCTGGT CCTGGCGAG CCCCTGGT CCAAGGGGG
-30 LeuglyAs nserylAsp ArgAlaProA rgProProAs pglyArgGly ArgValArgP roArgThrG1 nAspGlyVal GlyAsnHist hrMetAlaArg

201 GATCCCCAG ACCCTAAAGT TCGTGTGCTG CATCGTGGCG GTCTGTGCTGC CAGTCCTAGC TTRACTGTGCC ACCACTGTGCC ACCAGGAGGA AGTCCCCAG
CTAGGGTTC TGGGATTCA ACCGAGGAGCA GTAGCAGCG CAGGACGAG GTcAGGATCG AATGAGACGG TGGTACGGG CCGTCCTCTC TCAAGGGGT
-4 IleProLys ThrLeuLysP hevalVala lIleValAla ValleLeuP roValleAla atySerAla ThrThrAla rgLglngL uValProGln

301 CAGACAGTGG CCCAACAGCA ACAGAGGCC ACAGCTCAAGG GGGAGGACTG TCCAGGAGGA TCTCATAGAT CAGAACATAC TGGAGCCGT AACCCGTGCA
GTCGTCAAC GGGGTGTGT TGTCTCCGT TCGAAGTCC CCGTCCCTCAC AGGTGTCCT AGACTATCTA GTCTGTATC ACCTGGACA TTGGGACAGT
-37 GlnThrVala LaProGlnL nglnArgHis serPhelysG lygluglucy sProAlaGly serHisArgS ergluHist rGlyAlaCys AsnProCysthr

401 CAGAGGGTGT GGATTACACC AACGCTTCCA ACAATGAAACC TTCTGGCTTC CCATGTACAG TTCTGAATC AGATCAAAA CATAAAAGTT CCTGGCACCAC
GTCTCCACA OCTAATGGG TGCGGAAGGT TCTTACTGG AAGAACGAAAG GGTACATGTC APACATTAG TCTAGTTTT GTATTTCAA GGACGTGGTA
-71 GluglyVa IasptyrThr AsnAlaSerA snAsnLgUp serCysPhe ProCysthr alcySlyse raspglnLys HisLyssers erCysthrMet

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-104 ThrArgAsp ThrValcysG IncyslysG1 uglyThrPhe ArgAsnGlu snserProG1 umetCysArg LysCysSerA rgCysProse rglygluval

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GTTCACTCAT TAACATGGC GACCTACTA AACCTCTTAA ACCACGGTTA CGGTGACACC TTTGGGGTGC ACGACTTCTC TGTACTTGT
-137 GlnValSerA sncysthrse rTrpAspAsp IleGlnCysV algluglup eGlyAlaAsn AlaThrValG luthrProAl aAlaGluGlu ThrMetAsnThr

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GGTCGGCCCG CTGAGGACGG GTGAGGACAC TTCTCTGTTA CTTGTGGTGC GGCCTCTGAG GACGGGGTGC ACGACTTCTC TGTACTTGT
-171 SerProG1 yThrProAl proAlaAlaLugluthrme tasnThrSer ProGlyThrP roAlaProAl aAlaGluGlu ThrMetThrT hrSerProG1 yThrProAla

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-204 ThrProAla proAlaAlaLugluthrme tThrThrSer ProGlyThrP roAlaProAl aAlaGluGlu ThrMetThrT hrSerProG1 yThrProAla

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AGAAGAGTAA TGGAGACTAC GTGCTAGTAC CCCTAGTATC AAGATTAACA CGAAGACTAA CACAAACAA CTTCTGAAG TGACACCTTC TTAAGGAAG

237 serSerHist yrLeuSerCy sThrIleVal GlyIleIleV alleUileV aLLeUileV ValPheVal

1001 CCTTACCTGAA AGGTTCAAGGT AGGGGCTGGG TGAGGGGGG GGGGGCTGGA CACTCTCTGC CTCAGCTCTC TCTGCTGTGT TCCCACAGAC AGAACGCCT
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Apo2DcR	1	-----MARIPIKTLKFVV
DR4	51	GRGALPTSMQHGPSARARAGRAPGP P PAREASPRLRVHKT
		KFVVVGVD
Apo2	41	VVAAVLLILVSAESALITQODLAPOORAAPOOKRSSPSEG
Apo2DcR	13	LCPPGHHISED
DR4	101	VIVAVLIPVILAYSATTARQEEVPOOTVAPOOQRHSFKGEEC
		PAGSHRSER
Apo2	91	GRDCISCKYGDYSTHWNDLLFCERCTRCDSGEVELSPCTTTRNTVCOCE
Apo2DcR	63	TCACNPCTEGVGYTNASNNIFACIPLCTACKSDEEEERSPCTTTRNTACOCC
DR4	142	PGACNRCTEGVGYTNASNNIFACIPLCTACKSDEEEERSPCTTTRNTACOCC
		CRD1
Apo2	141	EGTFREEDSPEMCRKCRIGCPRGMVKVGDCTPWSDIECVHKE-----
Apo2DcR	113	EGTFERNENSPEMCRKCSR-CPSGEVQVSNCISWDDIQCVE-EFGANATVE
DR4	192	PGTFRNDNSAEMCRKCGSTGCPRGMVKVGDCTPWSDIECVHKE-----
		CRD2
Apo2	161	TPAAEETMNTSPGTPAPAAEETMNTSPGTPAPAAEETMTTSPGTPAPAAE
Apo2DcR		-----
DR4		-----
Apo2	183	-----SGIIIGVTVAAVVIIVAVFV-----
Apo2DcR	211	ETMTTSPGTPAPAAEETMTTSPGTPASSHYLSCTIVGIIVLVLLIVFV
DR4	234	-----SGNGHNIWVVELVVTLLVVPLILVAV-LIVC
Apo2	203	CKSLLKKVLPYLGICSGGGGDPERVDRSSQRPGAEDNVLNEIVSILQP
DR4	262	CCIGSGCGGDPKCMDRVCFWRLGLLRGPGAEDNAHNEILSNADSLSTFVS
Apo2	253	TQVPEQEMEVQEPAAEPITGVNMLSPGESEHLEPAEAERSORRRIIVPANE
DR4	312	-----EQOMESOEPADLTGVTVOQSPGEAQCLIGPAEAEGSORRRIIVPANG
Apo2	303	GDPTETELRCEDDFABLVPFDSWEPIMRKLGIMDNEIKVAKAEAAGH--R
DR4	358	ADPTETLMLFFDKTANIVPFDSDOLMRQLDLTKNEIDVVRAGTAGP--G
Apo3/DR3	338	VMDAVPARRWKEFVRTLGCREAEIEAVEVEI-GEF-R
TNFR1	322	VVENVPLRWKEFVERLIGLSDHEIDRLELQN-GCCLR
CD95	220	IAGVHTLSQVKGFVRKNGVNEAKIDEIKNDN-VQDTA
Apo2	351	DTLYTMLIKWVNKTGR-DASVHILLDAILETLGEFLAKOKIEDHLLSSGKF
DR4	406	DALYAMLMKVWNKTKGR-NASIHILLDALEMEERHAKETIQDILLVDSCKF
Apo3/DR3	374	DGQYEMLKRRRQQQP---AGLGAVYALLRMGLDCCVEDLRS
TNFR1	358	EAQYSMIAWRRTTPRREATLELIGRVLIRDMDLLGCLEDIEE
CD95	256	EUKVQLLERNWHQLHGKKEAY-DTIIKDIKKANLCTLAEKIQT
Apo2	400	MYLEGNA DALS
DR4	455	I Y LEDGTGS AVSLE

Fig. 2

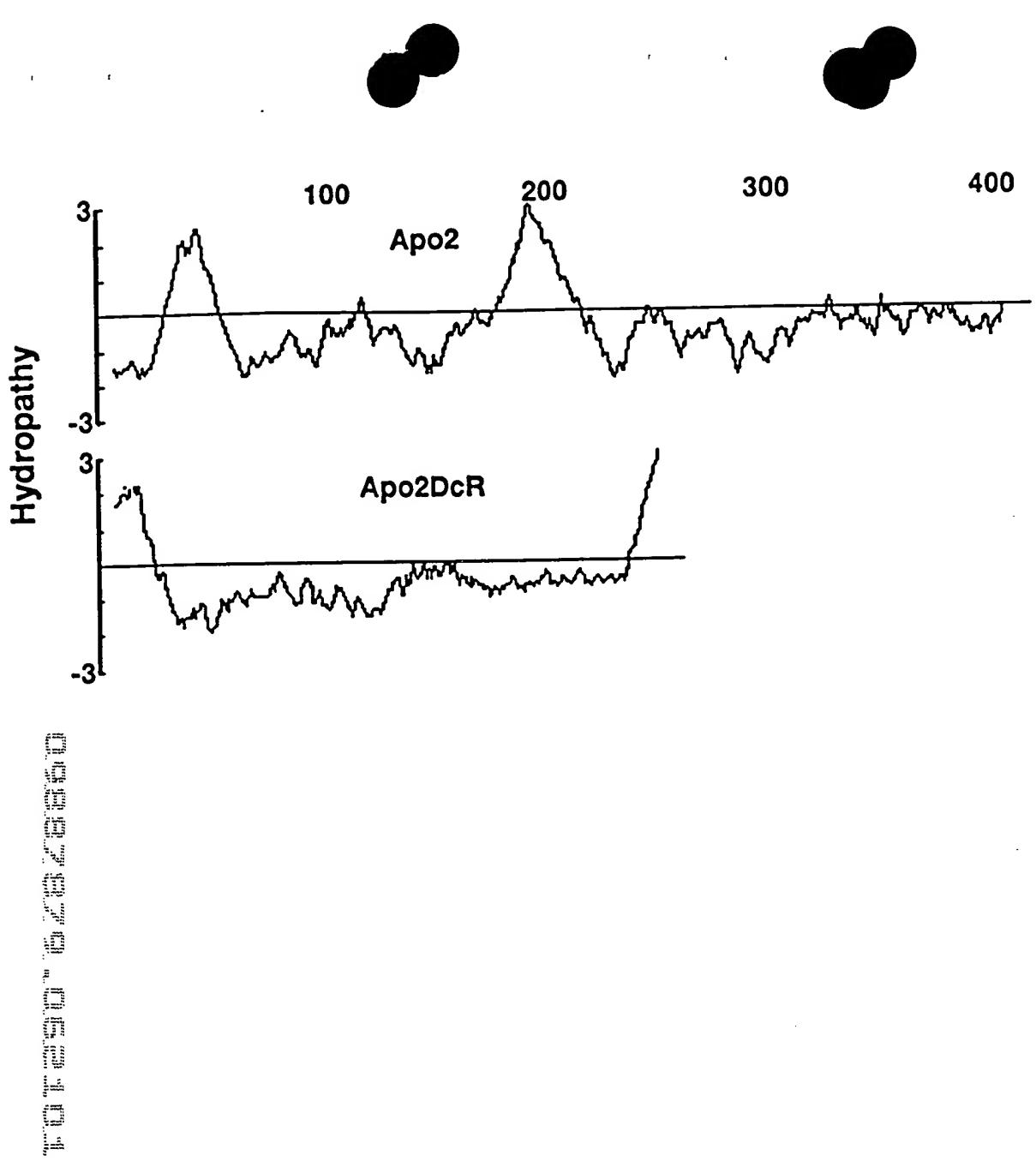


Figure 3

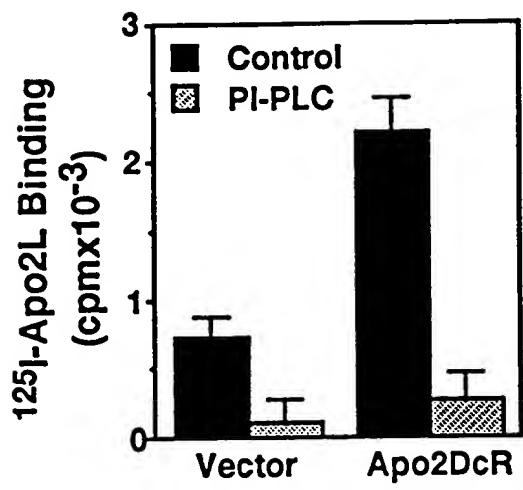


Figure 4

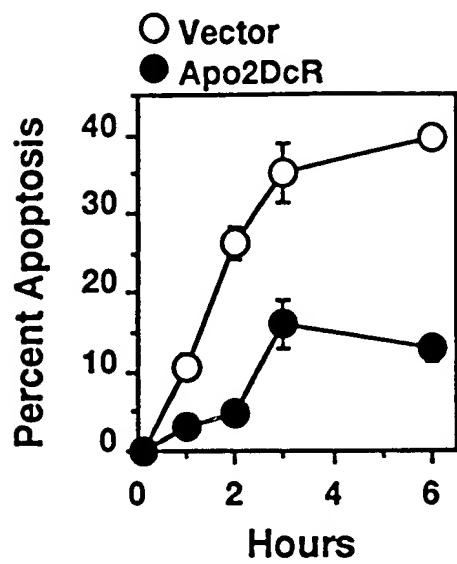


Figure 5

0 0 0 0 0 0 0 0 0 0



Figure 6

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Fig. 7A

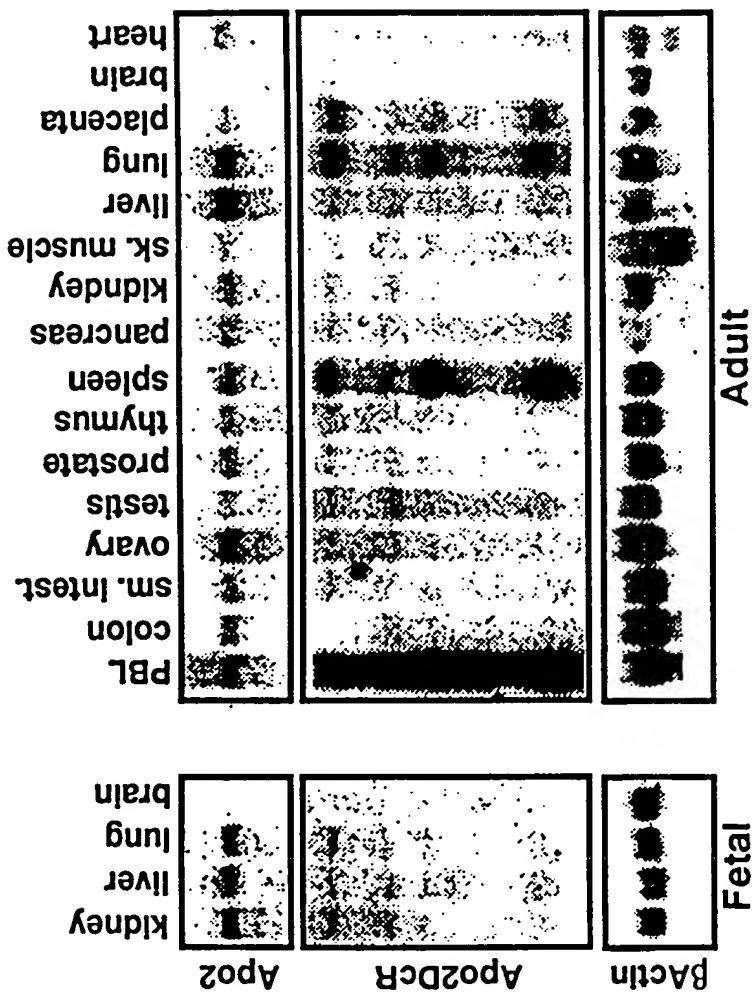


Fig. 7B

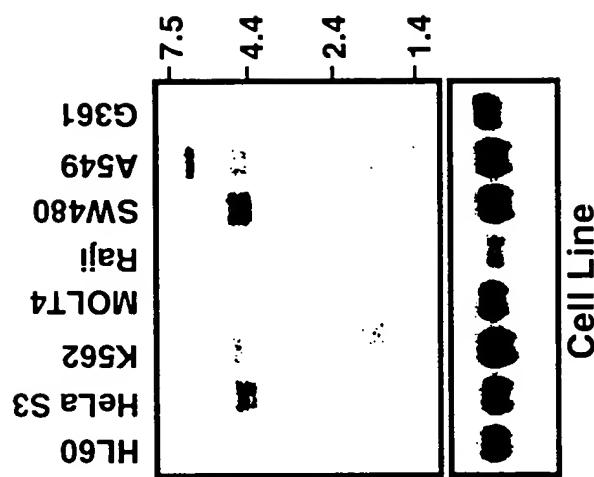


Fig. 8

Fig. 8
Protein sequence alignment

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GGTGGCGAG GCGTATTAG TCGTGGCGG GCCTCTGGG CGCTTAGAGA CGGGGTGTT TTATGTGGT GCTACGGGCT AGATGAAATT CCCGACTTTG

101 CCACGGGCT GAGAGACTAT AAGAGCGTTCC CCTAACCGCCA TGGAACACAACG GGGCACAGAAC GCCCCGGCC CTTGGGGGG CCGGAAAAGG CACGGCCAG
GGTGGCGGA CTCTCTGATA TTCTCGCAAG GGATGGGGT ACCTTGTTG CCGTGTCTG CGGGGGGGC GAAGCCCCCG GGCTTTCC GTGGCGGGGT
1 MetGluglnar gglylnasn AlaProAla laserGlyAl aArgLysArg HisGlyProGly

201 GACCAGGGG GCCAGGGGG CGCTCGGGT CGCTCGGT CCCAACAGCC CTTGCTGCTG TTGTCGCCCG GTGCCACCTGG ACCACATATC
CTGGTCCCT CGCGCCCT CGCTCGGGAC CCGAGGGCCA GGGGTTCTGG AACACCGAGC CCAGGGCCG CCAGGACGAC AACACAGTC GACTCAG;
22 ProArg1 uAlaArgGly AlaArgProGly LeuValLeu 1ProLySthr LeuValLeu alvalAla1 avalleLeu LeuValsera lagluser...

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AGACTAGTGG GTGTTCTGG ATCGAGGGGT CGTCTCTGC CGGGGTGTT TTTTCTCCAG GTCGGGGAGT CTCCCTAACM CAGGTGGACC TGTGGTATAG
55 LeuIlethr GlnglnasPL euAlaProG1 nglnargala AlaProGln InLysArgSe rserProser GluglyLeuc ysProProG1 yHishisile

401 TCAGGAAGCC GTAGAGATTG CATCTCCTGC AAATATGGAC AGGACTATAG CACTACTGG AATGACCTCC TTTTCTGCTT GGCCTGCCACC AGGTGTGATT
AGTCTTCTGC CATCTCTAAC GTAGAGGACG TTATACCTG TCCTGATATC GTGAGTGACC TTACTGGAGG AAAAGACGAA CGCGACGTGG TCCACACTAA
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501 CAGGTGAAGT CGACCTAACCA CGACCAAGAA CACAGTGTGT CAGTCCGAAG AAGGCACCTT CCGGGAAAGAA GATTCTCCCTG AGATGTGCG
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122 GlyGluva Igluleuser ProCysThr hrThrArgas nThrValCys GlncysGlug luglyThrPh eArgGluGlu AspSerProG lumetCysArg

601 GAAGTGGCCG ACAGGGGAT GGTCAGGGTC GGTGATTGTA CACCCTGGAG TGACATCGAA TGTGTCACAA AAGAA'TCAGG CATCATCATA
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155 LysCysArg ThrglycysP roArgGlyme tvallysval GlyAspCyst hrProTrpse raspileGlu CysValHisL ysGluSerG1 yllellle

701 GGAGTCACAG TTGCAAGGGT AGTCTTGATT GTGGCTGTGTT GTCTTACTG TGGAAAGAAG CCTCTCCTTA CCTGAAAGGC ATCTGCT
CCTCACTGTC AACGTGGCA TCAGAACTAA CACCGACACA AACAAACGTT CAGAAATGAC ACCTTCTTC AGGAAGGAAAT GGACTTCCG TAGACGAGTC
188 GlyValThrV alAlaAlava lvalleuLe ValAlaValP heValCysly sserLeuLeu TrpLyslyv alleuProty rleuLysGly lleCysSerGly

801 GTGGTGGTGG GGACCCCTGAG CGTGTGGACA GAAGCTCACA ACGACCTGG GCTGAGGACA ATGTCTCAA TGACATGTC AGTATCTTGC AGCCCACCCA
CACCAACCACC CCTGGGACTC GCACACCTGT TTCTCGAGTGT CGACTCCTGT TACAGGAGT ACTCTAGCAC TCATAGAACG TCGGGTGGGT
222 GlyGlyG1 YasProGlu ArgValAspa rgSerSerG1 nArgProGly AlaGluAspa snValLeuAs nGluIleVal serileLeuG lnProThrGln

901 GGTCCTGAG CAGGAATGG AAGTCCAGGA CCCACAGAG CCAACAGGTG TCAACATGTT GTCCCCGGG GAGTCAGAGC ATCTGCTGGA ACCGGCAGAA
CCAGGGACTC GTCCCTTACCG TTCAGGGTCTC CGTGTGTCAC AGTGTACAA CAGGGGGCC CTCAGTCTCG TAGACGACCT TGGGGTGTCT
255 ValProGlu GlnglumetG luvAlgLng1 uProAlaGlu ProThrGlyV alAsnMetLe userProGly GluLysLuh isLeuLeuG1 uprolaGlu

1001 GCTGAAAGGT CTCAGAGGAG GAGGTGGCTG GTTCCACTGAG TCCCACTGAG TCAACATGTT GTCCCCGGG GAGTCAGAGC ATCTGCTGGA ACCGGCAGAA
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288 AlaGluArgS ergInArgar gArgLeuLeu ValProAlaA snglyAs pProThrGlu ThrLeuArgG InCysPheAs paspPheAla AspLeuValPro

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 1102 GCAAACCTGAG GACCCTGGC GAGTACTCCT TCAACCCGCA GTACCTGTTA CTCTATTTCC ACCGATTTCG ACTCCGTCGC CGGTGTCCT TGTTGAACAT
 1103 322 Pheaspse RtrpgluPro Leumetargl ysleuglyle umetasparn GluileLysV alalalysal aglualaala GlyHisArga sptheleutyr.

 1201 CACGATGCTG ATAAAGTGGG TCAACAAAC CGGGCGAGAT GCCTCTGTC ACACCCCTGCT CGATGCCCTG GAGAGGACT TGCCAAAGCAG
 1202 GTGCTACGAC TATTTCACCC AGTTGTTTG GCCCGCTCTA CGGAGACAGG TGTGGACGA CCTACCGAAC CTCTGCGACC CTCTCTGCA ACGCTGTC
 1203 355 Thrmelteu IleLystrpv alasnLysth rgyaryasp Alaservalh isthrLeule uaspalaLeu GluthrLeug lygluargle ualalysGln

 1301 AAGATGAGG ACCACTGTT GAGCTCTGG AAGTTCAATGT ATCTAGAAGG TAATGCCAGAC TCTAAGTGTG ATTCTCTCTCA GGAAGTGAGA
 1302 TTCTAATCTC TGGTGAACAA CTCGAGACCT TTCAAGTACA TAGATCTTCC ATTACGTCTG AGACGGAAACA GGATTCACAC TAAGAGAAGT CCTTCACTCT
 1303 388 LysileGluA sphisLeule usersergLy LysPheMetT yrIeuglugl YasnAlaAsp SerAlaXqS eroC*

 1401 CCTTCCTGG TTACCTTT TTCTGGAAAA AGCCCAACTG GACTCCAGTC AGTAGGAAAG TGCCACAATT GTCACATGAC CGGTACTGGA AGAAACTCTC
 1402 GGAAGGGACC AAATGGAAA AAGACCTTT TCGGGTTGAC CTGAGGTCACT TCATCCTTC ACGGTGTTA CAGTGTACTG GCCATGACCT TCTTGAGA

 1501 CCATCCAACA TCACCCAGTG GATGGAACAT CCTGTAACCT TTCACTGCACT TTGGCAATT TTTATAAGC TGAATGTGAT ATAAGGACA CTATGGAAT
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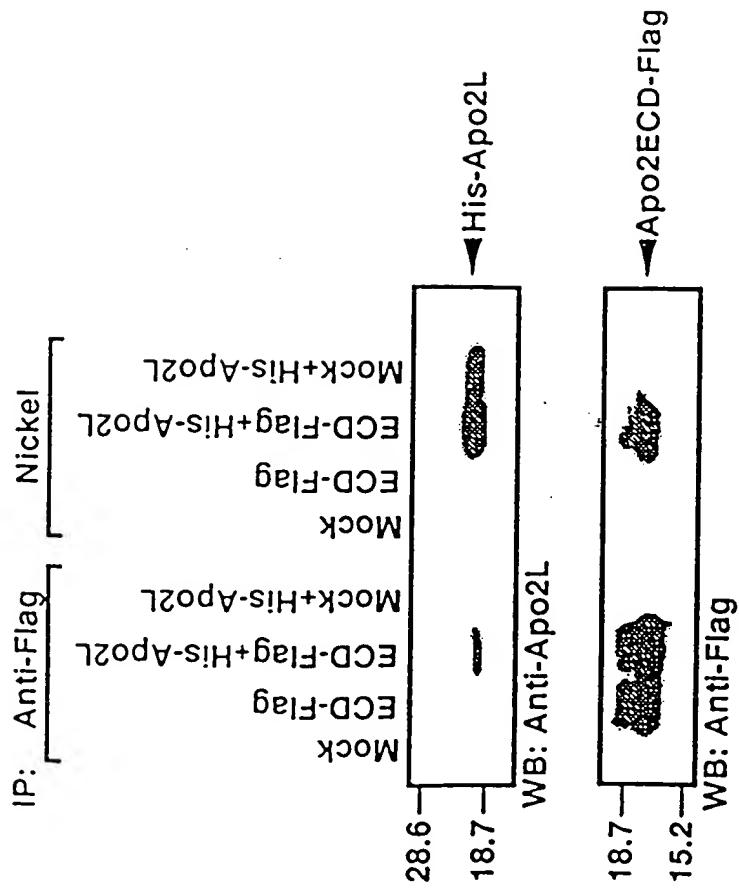
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Fig. 8 (cont.)

Fig. 9

1 MEQRGONAPAAASGARKRHC~~P~~GPREARGARPGLRVPKTLVLUVAAVL~~L~~VSAESALITQQD
61 LAPQORAAPQQKRS~~S~~PPSEG~~L~~CPPGH~~H~~I~~S~~EDGRDCIS~~C~~KYQDYSTHWNDL~~L~~FCLR~~C~~TRCD
121 SGEVEILSP~~C~~TITRNTV~~C~~QE~~E~~GT~~F~~REEDSP~~E~~MCRKCRTG~~C~~PRGMVKVG~~D~~CTPWSDIE~~C~~VH
181 KE~~S~~G~~I~~IIIGVTVAAVLLIVAFVU~~C~~KSLLM~~K~~KVLPYLKGICSGGGDPERVDRSSQRGAED
241 NVLNEIVSILQOPTQVPEQEMEVQEPAA~~P~~PTGVNM~~L~~SPGES~~E~~SHLLEPAEAERSQRRLLVPA
301 NEGDPTE~~T~~LRQC~~F~~DDFADLVPSWEPLMRKLGLMDNEIKVAKAAAGHRDTLYTMLLIKW
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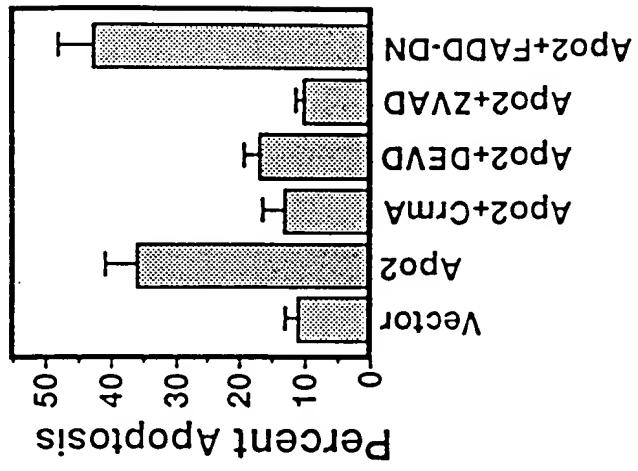
SCANNED, #12
Fig. 10



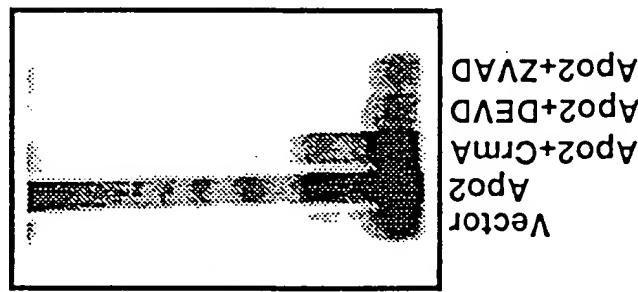
SCANNED, #12

Fig. 11

III C



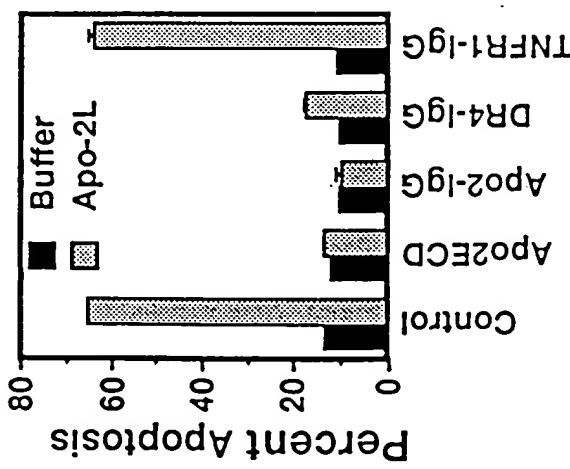
III B



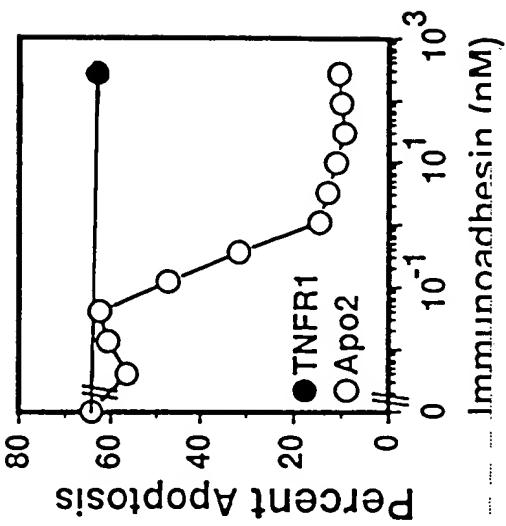
III A



III D

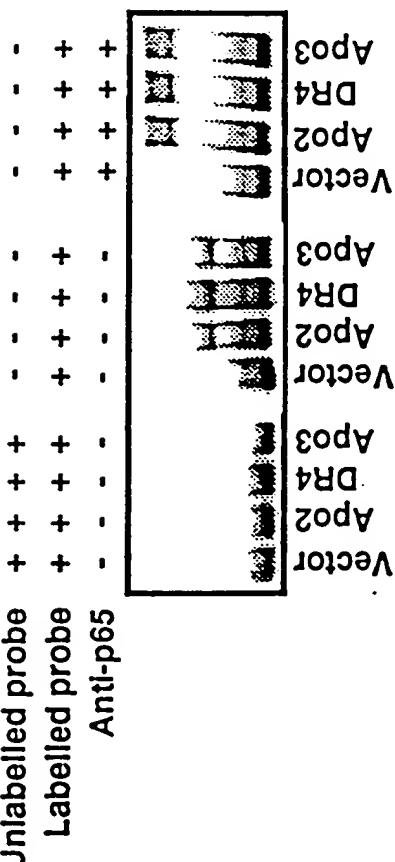


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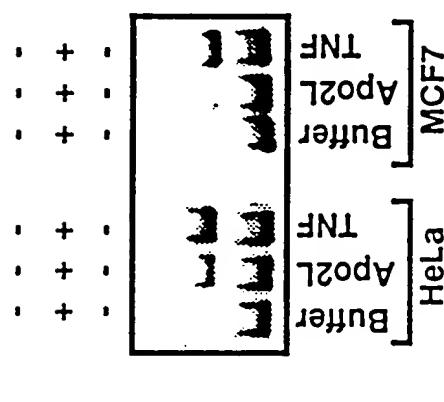


SCANNED, #12

12A



12B



12C

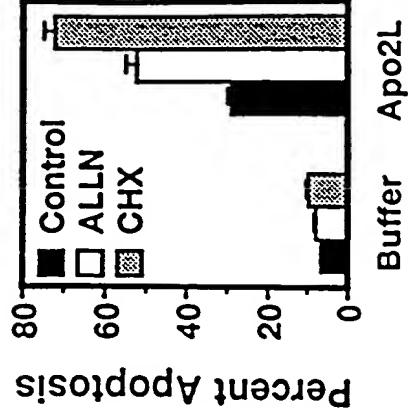
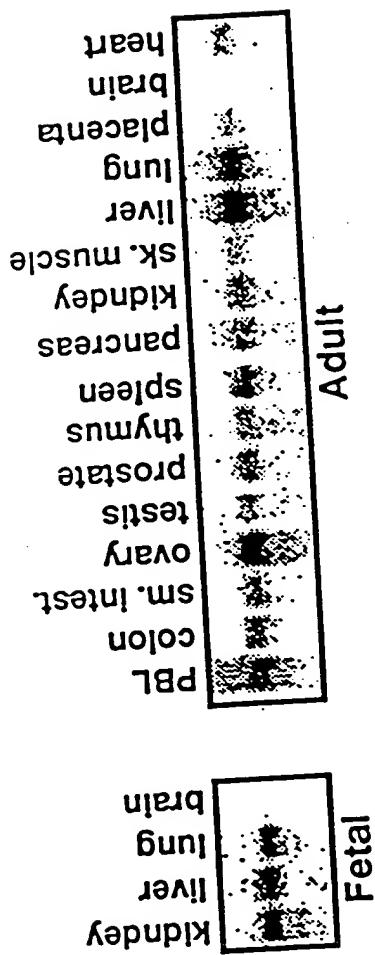


Fig. 12

Fig. 13



1C5.24.1
4G3.9.9
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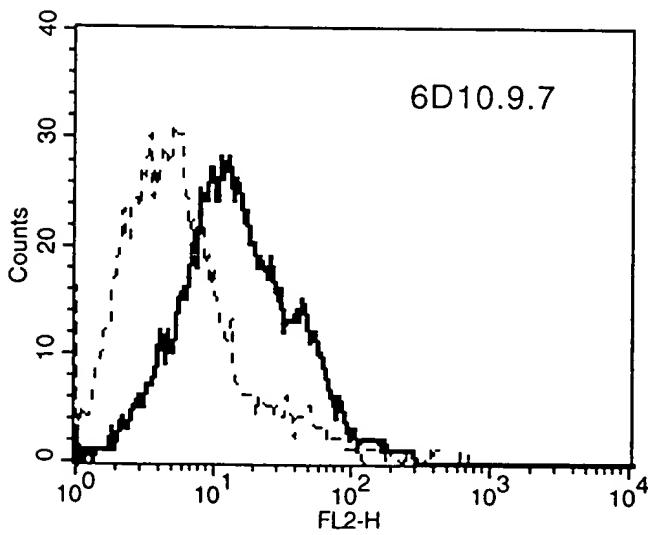
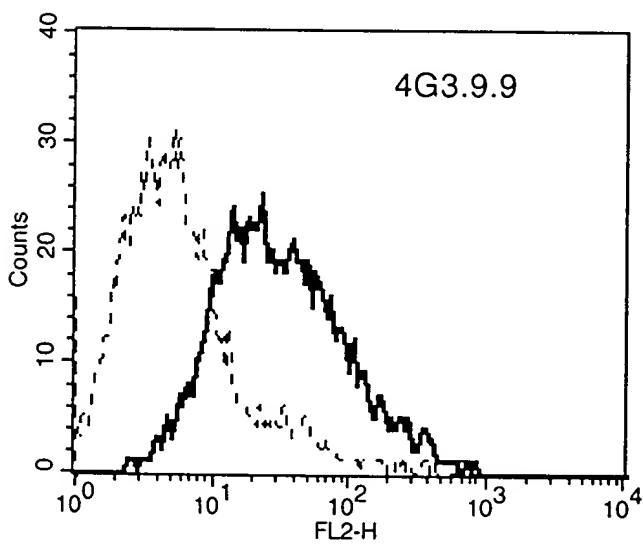
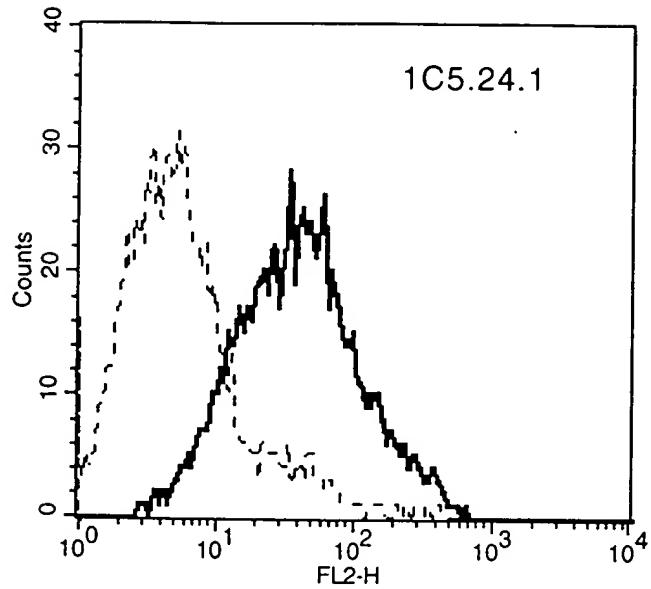


Fig. 14

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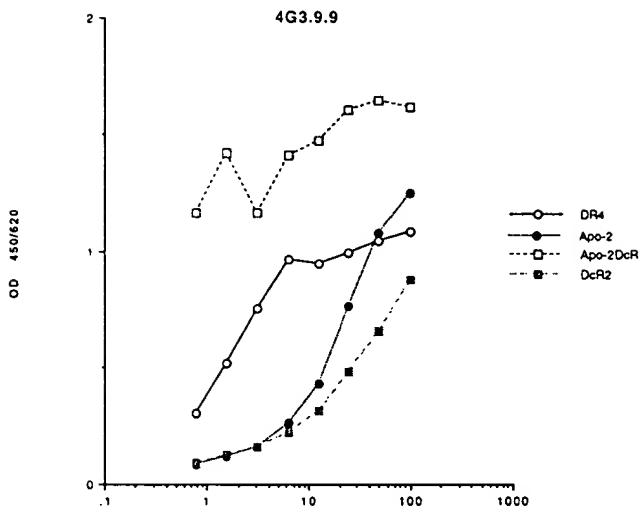
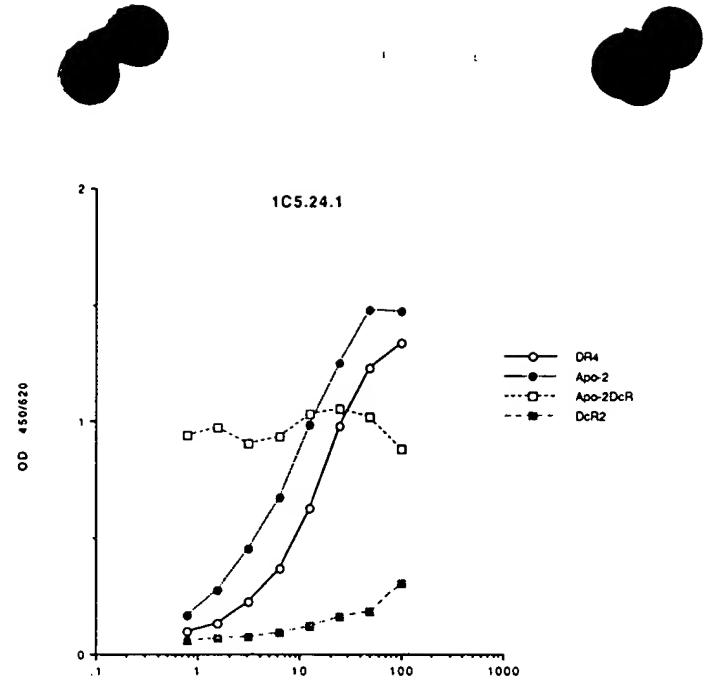
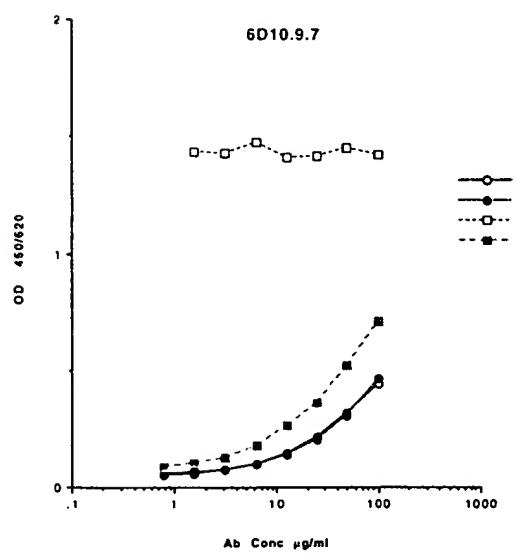


Fig. 15



Summary of mAbs to DcR1

mAbs	ISOTYPE	FACS (HUMEC)	DR4	Cross reactivity		
				$\Delta_{\text{apo}-2}$	$\Delta_{\text{apo}-2\alpha R}$	DcR2
1C5.24.1	IgG1	+	++	+++	+++	-
4G3.9.9	IgG1	+	++	+	+++	+/-
6D10.9.7	IgG2b	+	-	-	+++	+/-

Percent Cross reactivity was determined by comparing the binding capacity to $\Delta_{\text{apo}-2\alpha R}$ at 10 ug/ml of mAbs in ELISA. ++: >75% , +: 25-75% , +/-: 10-25% , -: <10% .

Fig. 16

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